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**IN THE UNITED STATES PATENT AND TRADEMARK OFFICE**

Applicants: Jeffrey Tze Fei Wong et al.

Serial No.: 09/827,255

Title: COMPOSITIONS AND METHODS FOR THE TARGETED DELIVERY OF AGENTS TO TREAT LIVER CANCER

Filing Date: May 4, 2001

Art Unit: 1636

Examiner: M. Schmidt

Mail Stop RCE  
Commissioner for Patents  
P.O. Box 1450  
Alexandria, VA 22313-1450

**REQUEST FOR CONTINUED EXAMINATION**

Dear Sir:

In response to the Advisory Action of June 12, 2003, please amend the above-identified application as follows. The final office action was mailed on February 6, 2003. Response was due May 6, 2003 and one-month extension of time was paid with the response filed on June 5, 2003, making the response timely filed on June 5, 2003. Applicants hereby enclose an additional two-month petition for extension of time extending the period for response to August 6, 2003, making this response timely filed.

Amendments to the Claims are reflected in the listing of claims that begins on page 2 of this filing.

Remarks / Arguments begin on page 4 of this filing.

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## AMENDMENTS

This listing of claims will replace all prior versions, and listings, of claims in the application. Support for the amendments may be found throughout the specification, including the claims as originally filed.

1. (currently amended) A composition for the targeted delivery of an therapeutic active agent to a tissue expressing asialoglycoprotein receptors comprising an effective amount of the agent encapsulated in a liposome having a molar ratio PC:Chol:PS of 11:42:0.025 coupled to desialylated glycoprotein- $\alpha$ 1.
2. (currently amended) The composition of claim 1, wherein the therapeutic agent is a drug or a polynucleotide.
3. (currently amended) The composition of claim 2, wherein the polynucleotide is cDNA encoding a therapeutic protein, a ribozyme, and antisense DNA.
4. (currently amended) The composition of claim 1, wherein the therapeutic agent is selected from the group consisting of a cytotoxic drugs, a protein.
5. (original) The composition of claim 4, wherein the cytotoxic drug is selected from the group consisting of doxorubicin, vincristine, daunorubicin, and amphiphatic amines.
6. (previously amended) The composition of claim 1, wherein the desialylated glycoprotein- $\alpha$ 1 is coupled to the liposome by an avidinbiotin or thiol-maleamide linkages.
7. (currently amended) A method for targeted delivery of a therapeutic active agent to a tissue expressing asialoglycoprotein receptors comprising delivery to the tissue an effective amount of the composition of any of claims 1 to 6.
8. Canceled

9. (currently amended) A composition for the targeted delivery of a therapeutic  
active agent to a tissue expressing asialoglycoproteing receptors comprising an effective amount  
of a doxorubicin encapsulated in a liposome having a molar ratio PC:Chol:PS of 11:0.025  
coupled to desialytated glycoprotein-  $\alpha$  1 by anavidin-biotin linkage.

10. Withdrawn